

CLAIMS

1. A method of producing hybrid proteins from a hybrid gene cDNA library comprising:

5 providing a purified sample of a vector comprising a DNA molecule having at least one selectable marker sequence and a sequence encoding a hybrid protein region, wherein the hybrid protein region comprises:

a regulatable DNA sequence;

10 a multiple cloning site immediately 3' to the regulatable DNA sequence, wherein the multiple cloning site does not encode a translational termination sequence; and

15 a DNA sequence encoding at least one common peptide placed 3' to the multiple cloning site, wherein the common peptide encoding sequence does not contain a translation initiation codon;

isolating a mRNA template population of interest;

20 synthesizing a cDNA population from the mRNA template population using random sequence oligonucleotide primers;

adding cloning linkers to the cDNA population;

cleaving the vectors at the multiple cloning site;

25 inserting the cDNA population molecules into the cleaved vectors, to create a hybrid gene cDNA library;

transforming bacterial cells with the hybrid gene cDNA library and selecting transformed cells;

purifying the hybrid gene cDNA library from the transformed bacterial cells;

transforming yeast cells with the hybrid gene cDNA library and selecting transformed cells; and

5 allowing transformed yeast cells to produce a hybrid protein.

2. The method of claim 1, wherein the bacterial cells transformed with the hybrid gene cDNA library are

10 *E. coli* cells.

3. The method of claim 1, wherein the vector encodes a common peptide sequence comprising six successive histidine residues and the hybrid protein is

15 purified from the yeast cells using affinity purification.

4. The method of Claim 1, wherein the hybrid protein region further comprises a transcription

20 termination sequence placed immediately 3' to the common peptide encoding sequence.

5. A hybrid protein production method comprising:  
isolating an mRNA template population;  
synthesizing a cDNA population from the mRNA  
template population using random sequence oligonucleotide  
5 primers;  
cleaving vectors at a multiple cloning site;  
inserting members of the cDNA population into the  
cleaved vectors, to create a hybrid gene cDNA library;  
and  
10 expressing a hybrid protein from the hybrid gene  
cDNA library.

6. The method of Claim 5, wherein the vectors  
further comprise a DNA molecule having at least one  
15 selectable marker sequence and a hybrid protein region  
sequence.

7. The method of Claim 6, wherein the hybrid  
protein region sequence further comprises:  
20 a regulatable DNA sequence;  
a multiple cloning site lacking a translation  
termination sequence placed immediately 3' to the  
regulatable DNA sequence; and  
at least one common peptide encoding sequence  
25 lacking a translation initiation codon placed 3' to the  
multiple cloning site.

8. The method of Claim 7, wherein the hybrid protein region sequence further comprises a transcription termination sequence placed immediately 3' to the common 5 peptide encoding sequence.

9. The method of Claim 5, further comprising:  
transforming bacterial cells with the hybrid gene cDNA library and selecting transformed cells;  
10 purifying the hybrid gene cDNA library from the transformed bacterial cells;  
transforming yeast cells with the hybrid gene cDNA library and selecting transformed cells; and  
expressing the hybrid protein in the transformed 15 yeast cells.

10. The method of Claim 9, wherein the bacterial cells comprise *E.coli*.

20 11. The method of Claim 5, wherein the vectors encode a common peptide sequence having six successive histidine residues and further comprising purifying the hybrid protein using affinity purification.

12. A hybrid protein production method comprising:  
isolating an mRNA template population;  
synthesizing a cDNA population from the mRNA  
template population;

5        cleaving vectors at a multiple cloning site, wherein  
the vectors include a DNA molecule having at least one  
selectable marker sequence and a hybrid protein region  
sequence including:  
            a regulatable DNA sequence;

10        a multiple cloning site lacking a translation  
termination sequence placed immediately 3' to the  
regulatable DNA sequence; and  
            at least one common peptide encoding sequence  
lacking a translation initiation codon placed 3' to the  
15        multiple cloning site;

            inserting members of the cDNA population into the  
cleaved vectors, to create a hybrid gene cDNA library;  
and  
            expressing a hybrid protein from the hybrid gene  
20        cDNA library.

13. The method of Claim 12, wherein synthesizing  
the cDNA population comprising using random sequence  
oligonucleotide primers.

14. The method of Claim 12, wherein the hybrid protein region sequence further comprises a transcription termination sequence placed immediately 3' to the common peptide encoding sequence.

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15. The method of Claim 12, further comprising:  
transforming bacterial cells with the hybrid gene cDNA library and selecting transformed cells;  
purifying the hybrid gene cDNA library from the  
10 transformed bacterial cells;  
transforming yeast cells with the hybrid gene cDNA library and selecting transformed cells; and  
expressing the hybrid protein in the transformed yeast cells.

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16. The method of Claim 15, wherein the bacterial cells comprise *E.coli*.

17. The method of Claim 12, wherein the vector  
20 encodes a common peptide sequence having six successive histidine residues and further comprising purifying the hybrid protein using affinity purification.

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